

Fig. S1. Inhibition of cell viability by trastuzumab, as measured by cell counting kit-8
assay, occurred in a dose-dependent manner 4 days following treatment with
trastuzumab at five different concentrations. Data are presented as mean ± standard
deviation (SD). The data shown are representative of three independent experiments.
\*\*\*p < 0.001, \*\*p < 0.01.</li>



8 Fig. S2. Trastuzumab ( $10 \mu g/mL$ ) inhibited cell viability in a time-dependent manner.

- 9 Data are presented as mean  $\pm$  standard deviation (SD). The data shown are
- 10 representative of three independent experiments. \*\*p < 0.01.



Fig. S3. YAP knockdown. (A) YAP protein and N-YAP expression were detected,
using western blotting, in BT474-TS cells. (B) YAP mRNA levels detected, by

14 RT-PCR, in BT474-TS cells. Data are presented as mean  $\pm$  standard deviation (SD).

15 The data shown are representative results of three independent experiments. \*\*p <

16 0.01.



**Fig. S4.** Study hypothesis: AKT attenuates p73-mediated apoptosis by

<sup>19</sup> phosphorylating YAP.



21 Fig. S5. Combination treatment of AKT inhibitor (GSK) and trastuzumab of

22 YAP-over BT474-TR cells. (A) Trastuzumab plus GSK increased apoptosis rate of

23 BT474-TR cells. (B) Expression of apoptotic proteins was examined using western

24 blotting. (C) Trastuzumab plus GSK strongly affected the viability of BT474-TR cells

after YAP-overexpression. Data are presented as mean  $\pm$  standard deviation (SD). The

data shown are representative of three independent experiments. \*p < 0.05.



Fig. S6. p73-overexpression altered trastuzumab treatment effects in YAP-silenced 28 BT474-TS cells. (A) p73 protein expression was detected, by western blotting, in 29 YAP-silenced BT474-TS cells. (B) p73-overexpression altered apoptosis, measured 30 31 by flow cytometry, following treatment of YAP-silenced BT474-TS cells. (C) Expression of apoptotic proteins was examined by western blotting. (D) 32 p73-overexpression decreased the viability of YAP-silenced BT474-TS cells after 33 34 trastuzumab treatment; Data are presented as mean  $\pm$  standard deviation (SD). The data shown are representative of three independent experiments. \*p < 0.05. 35



Fig. S7. YAP expression in breast cancer tissues. YAP protein expression was detected, by western blot, in pCR and non-pCR breast cancer tissues. Data are presented as mean  $\pm$  standard deviation (SD). The data shown are representative of three independent experiments. \*p < 0.05.

				YAP		
			No.	Negative	Positive	<i>p</i> -value
				No. (%)	No. (%)	
TAC		negative	9	7 (77.8)	2 (22.2)	
	p73	positive	14	5 (35.7)	9 (64.3)	0.049
		negative	4	1 (25.0)	3 (75.0)	
	AKT	positive	19	11 (60.0)	8 (40.0)	0.291
		negative	7	1 (14.3)	6 (85.7)	
	p-AKT	positive	16	11 (68.8)	5 (31.2)	0.016
ТСЬН		negative	6	6 (100)	0 (0)	
	p73	positive	8	1 (12.5)	7 (87.5)	0.001
			2		2 (100)	
		negative	3	0 (0)	3 (100)	
	AKT	positive	11	7 (63.6)	4 (36.4)	0.051
		negative	7	0 (0)	7 (100)	
	p-AKT	positive	7	7 (100)	0 (0)	< 0.001

Table S1. Association between YAP and p73, AKT, and p-AKT in pre-treatment biopsy tissues

				YAP		
			No.	Negative	Positive	<i>p</i> -value
				No. (%)	No. (%)	
TAC		negative	6	5 (83.3)	1 (16.7)	
	p73	positive	14	6 (42.9)	8 (57.1)	0.095
		negative	10	3 (30 0)	7 (70.0)	
		inegative	10	3(50.0)	7(70.0)	0 122
	AKT	positive	13	8 (61.5)	5 (38.5)	0.133
		negative	12	3 (25.0)	9 (75.0)	
	p-AKT	positive	8	8 (100)	0 (0)	0.001
					0 (0)	
ТСЬН		negative	4	4 (100)	0 (0)	
	p73	positive	6	0 (0)	6 (100)	0.002
		negative	3	0 (0)	3 (100)	
	AKT	positive	7	4 (57.1)	3 (42.9)	0.091
		negative	6	0 (0)	6 (100)	
	p-AKT	positive	4	4 (100)	0 (0)	0.002

**Table S2.** Association between YAP and p73, AKT, and p-AKT in post-treatment surgical tissues